

AMENDMENTS TO THE CLAIMS

Listing of Claims

This listing of the claims will replace all prior versions, and listings, of claims in this application.

1 – 59. **(Cancelled)**

60. **(Currently Amended)** A method of identifying compounds that bind to a leukotriene A₄ (LTA₄) hydrolase comprising the amino acid sequence of SEQ ID NO:1, the method comprising the steps of:

(a) crystallizing a purified LTA₄ hydrolase in the presence of bestatin to form a co-crystal of ~~[[an]] LTA₄ hydrolase and bestatin~~ crystal, wherein crystallization is performed ~~[[as]]~~ by liquid- liquid diffusion in a capillary using equal volumes of a buffer: enzyme solution consisting of:

i) a buffer solution consisting of 28% PEG8000, 0.1 M Na-acetate, 0.1 M imidazole at a pH of 6.8 and with 5 mM YbCl₃ as an additive; and

ii) an enzyme solution consisting of 5 mg/ml LTA₄ hydrolase comprising the amino acid sequence of SEQ ID NO:1 in 10 mM Tris-HCl at a pH of 8, supplemented with 1 mM bestatin;

wherein the crystallization results in a LTA₄ hydrolase crystal having the space group P2₁2₁2 and the unit cell dimensions a=67.59 Å, b=133.51 Å, and c=83.40 Å and $\alpha=\beta=\gamma=90^\circ$;

(b) determining the atomic coordinates of said LTA₄ hydrolase crystal; and

(c) screening the atomic coordinates of a set of candidate compounds against the atomic coordinates of said LTA₄ hydrolase crystal obtained in step a) to identify compounds that bind to the LTA₄ hydrolase;

~~wherein the crystallization results in a LTA₄ hydrolase crystal having the space group P2₁2₁2 and the unit cell dimensions a=67.59 Å, b=133.51 Å, and c=83.40 Å and wherein $\alpha=\beta=\gamma=90^\circ$.~~

61. **(Previously Presented)** The method of claim 60, wherein the LTA₄ hydrolase is purified by adsorption chromatography on hydroxyapatite and anion-exchange chromatography.

62-67. **(Cancelled)**

68. **(Previously Presented)** The method of claim 60, wherein the atomic coordinates of said LTA₄ hydrolase crystal correspond to the atomic coordinates defining atom 1 to atom 4876 as set forth in Table 9.

69. **(Cancelled)**

70. **(Currently Amended)** A method of designing an inhibitor or an agonist of LTA₄ hydrolase comprising the amino acid sequence of SEQ ID NO:1, the method comprising the steps of:

(a) crystallizing a purified LTA₄ hydrolase in the presence of bestatin to form a co-crystal of LTA₄ hydrolase and bestatin and thereafter determining its three dimensional conformational structure, wherein the crystallization is performed ~~[[as]]~~by liquid-liquid diffusion in a capillary using equal volumes of a buffer: enzyme solution consisting of:

i) a buffer solution consisting of 28% PEG8000, 0.1 M Na-acetate, 0.1 M imidazole at a pH of 6.8 and with 5 mM YbCl₃ as an additive; and

ii) an enzyme solution consisting of 5 mg/ml LTA₄ hydrolase comprising the amino acid sequence of SEQ ID NO:1 in 10 mM Tris-HCl at a pH of 8, supplemented with 1 mM bestatin;

wherein the crystallization results in a LTA₄ hydrolase crystal having the space group P2₁2₁2 and the unit cell dimensions a=67.59 Å, b=133.51 Å, and c=83.40 Å and α=β=γ=90°; and

iii) determining the atomic coordinates of said LTA₄ hydrolase crystal;

(b) identifying at least one potential inhibitor or agonist compound that is at least in part complementary to the LTA₄ hydrolase by the use of the atomic coordinates of the LTA₄ hydrolase crystal~~conformational structure of the crystal complex obtained in step a);~~

(c) soaking the co-crystallized LTA₄ hydrolase obtained in step a) with a solution of ~~a compound~~the potential inhibitor or agonist identified in step b) to obtain a complex of the crystal of said LTA₄ hydrolase and said potential inhibitor or agonist compound; and

(d) determining the atomic coordinates performing X ray crystallography of the crystal complex of said LTA₄ hydrolase and said inhibitor or agonist in step (c) compound to determine the structure thereof, thereby identifying the potential inhibitor or agonist compound as an inhibitor or agonist of LTA₄ hydrolase;

~~wherein the crystallization results in a LTA₄ hydrolase crystal having the space group P21212 and the unit cell dimensions a=67.59 Å, b=133.51 Å, and c=83.40 Å and wherein $\alpha=\beta=\gamma=90^\circ$.~~

71. **(Previously Presented)** The method of claim 70, wherein the LTA₄ hydrolase is purified by adsorption chromatography on hydroxyapatite and anion-exchange chromatography.

72-75. **(Cancelled)**

76. **(Currently Amended)** The method of claim 70, wherein the atomic coordinates of said LTA₄ hydrolase crystal in step (a) correspond to the atomic coordinates defining atom 1 to atom 4876 as set forth in Table 9.

77. **(Cancelled)**

78. **(Currently Amended)** The method of claim 70, further comprising the step of refining the structure of the inhibitor or agonist~~said compound obtained in step d) via computer modeling using data obtained in step d) via computer modeling and using this refined data~~from the X-ray crystallography in step d) and repeating steps b)-d).

79. **(Previously Presented)** The method of claim 70, wherein the complex obtained in step c) comprises bestatin.

80-86. **(Cancelled)**